Applicant's File Reference: 36119.136WO International Application No.: PCT/US05/08946

REPLACEMENT SHEETS – WITH CHANGES SHOWN What is claimed is:

- A method of screening for agents for treating asthma in a human,
 comprising:
 - (a) contacting a <u>mammalian</u> Gob-4 protein with a test agent thought to be effective in inhibiting the activity of said <u>mammalian</u> Gob-4 protein;
 - (b) determining if said test agent inhibits the activity of said mammalian Gob-4 protein, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises quantitating the amount of mucus produced by staining said mucus with a periodic-acid Schiff stain; and
 - (c) classifying said test agent as an agent for treating asthma if said test agent inhibits the activity of said <u>mammalian</u> Gob-4 protein.
- A method of screening for agents for treating asthma in a mammal,
 comprising:
 - (a) contacting a Gob-4 protein with a test agent thought to be effective in inhibiting the activity of said Gob-4 protein;
 - (b) determining if said test agent inhibits the activity of said Gob-4 protein; and
 - (c) <u>classifying said test agent as an agent for treating asthma if said test agent inhibits the activity of said Gob-4 protein.</u>
 - 3. The method of claim 2, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises determining the number of goblet cells that form from epithelial cells.
- 25 4. The method of claim 2, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises quantitating the amount of mucus produced.
 - 5. The method of claim 4, wherein said quantitating the amount of mucus produced comprises quantitating the amount of mucopolysaccharides in said mucus.

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- 6. The method of claim 2, wherein said Gob-4 protein has an amino acid sequence selected from the group consisting of SEQ ID NO:2 and SEQ ID NO:4.
- 7. The method of claim 2, wherein said Gob-4 protein has an amino acid sequence having at least about 70% identity to an amino acid sequence selected from the group consisting of SEQ ID NO:2 and SEQ ID NO:4.
- 8. <u>A method of screening for agents for treating asthma in a mammal, comprising:</u>
 - (a) contacting a nucleotide sequence encoding a reporter gene product operably linked to a Gob-4 protein promoter with a test agent thought to be effective in inhibiting production of a Gob-4 protein;
 - (b) <u>determining if said test agent inhibits production of said reporter gene</u> <u>product; and</u>
 - (c) <u>classifying said test agent as an agent for treating asthma if said test agent inhibits production of said reporter gene product.</u>
- 9. The method of claim 8, wherein determining if said test agent inhibits production of said Gob-4 protein comprises quantifying the amount or activity of said reporter gene product.
 - 10. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence selected from the group consisting of SEQ ID NO:5.
- 20 <u>The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence having at least about 80% identity to the nucleotide sequence selected from the group consisting of SEQ ID NO:5.</u>
 - 12. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence selected from the group consisting of SEQ ID NO:6.
- 13. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence having at least about 80% identity to the nucleotide sequence selected from the group consisting of SEQ ID NO:6.

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- 14. The method of claim 8, wherein said reporter gene product is selected from the group consisting of luciferase, β-galactosidase, chloramphenical acetyltransferase, β-glucuronidase, alkaline phosphatase, and green fluorescent protein.
- 5 15. A method for treating asthma, comprising administering to a mammal in need thereof a therapeutic amount of an agent that decreases the activity of a Gob-4 protein.
 - 16. A method for treating asthma, comprising administering to a mammal in need thereof a therapeutic amount of an agent that decreases the production of a Gob-4 protein.
 - 17. The method of claim 16, wherein said agent that decreases the production of said Gob-4 protein is a nucleic acid.
 - 18. The method of claim 17, wherein said nucleic acid is a ribonucleic acid.
- 19. The method of claim 18, wherein said ribonucleic acid has a nucleotide sequence that is complementary to a portion of the nucleotide sequence set forth in SEQ ID NO:1 or SEQ ID NO:3 encoding said acidic mammalian protein.
 - 20. The method of claim 18, wherein said ribonucleic acid is RNA interference.

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What is claimed is:

1. A method of screening for agents for treating asthma in a human, comprising:

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(a) contacting a mammalian_Gob-4 protein with a test agent thought to be effective in inhibiting the activity of said mammalian_Gob-4 protein;
(b) determining if said test agent inhibits the activity of said

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(b) determining if said test agent inhibits the activity of said mammalian_Gob-4 protein, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises quantitating the amount of mucus produced by staining said mucus with a periodic-acid Schiff stain; and

with a periodic-acid Schill Stain, and

(c) classifying said test agent as an agent for treating asthma if said test agent inhibits the activity of said mammalian_Gob-4 protein.

- 2. A method of screening for agents for treating asthma in a mammal, comprising:
 - (a) contacting a Gob-4 protein with a test agent thought to be effective in inhibiting the activity of said Gob-4 protein;
 - (b) determining if said test agent inhibits the activity of said Gob-4 protein; and

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- (c) classifying said test agent as an agent for treating asthma if said test agent inhibits the activity of said Gob-4 protein.
- 3. The method of claim 2, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises determining the number of goblet cells that form from epithelial cells.
- 4. The method of claim 2, wherein determining if said test agent inhibits the activity of said mammalian Gob-4 protein comprises quantitating the amount of mucus produced.
 - 5. The method of claim 4, wherein said quantitating the amount of mucus produced comprises quantitating the amount of mucopolysaccharides in said mucus.

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6. The method of claim 2, wherein said Gob-4 protein has an amino acid sequence selected from the group consisting of SEQ ID NO:2 and SEQ ID NO:4.

- 7. The method of claim 2, wherein said Gob-4 protein has an amino acid sequence having at least about 70% identity to an amino acid sequence selected from the group consisting of SEQ ID NO:2 and SEQ ID NO:4.
- 8. A method of screening for agents for treating asthma in a mammal, comprising:
 - (a) contacting a nucleotide sequence encoding a reporter gene product operably linked to a Gob-4 protein promoter with a test agent thought to be effective in inhibiting production of a Gob-4 protein;
 - (b) determining if said test agent inhibits production of said reporter gene product; and
 - (c) classifying said test agent as an agent for treating asthma if said test agent inhibits production of said reporter gene product.
- 9. The method of claim 8, wherein determining if said test agent inhibits production of said Gob-4 protein comprises quantifying the amount or activity of said reporter gene product.
 - 10. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence selected from the group consisting of SEQ ID NO:5.
 - 11. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence having at least about 80% identity to the nucleotide sequence selected from the group consisting of SEQ ID NO:5.
 - 12. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence selected from the group consisting of SEQ ID NO:6.
 - 13. The method of claim 8, wherein said Gob-4 protein promoter has a nucleotide sequence having at least about 80% identity to the nucleotide sequence selected from the group consisting of SEQ ID NO:6.

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- 14. The method of claim 8, wherein said reporter gene product is selected from the group consisting of luciferase, β -galactosidase, chloramphenical acetyltransferase, β -glucuronidase, alkaline phosphatase, and green fluorescent protein.
- 15. A method for treating asthma, comprising administering to a mammal in need thereof a therapeutic amount of an agent that decreases the activity of a Gob-4 protein.
- 16. A method for treating asthma, comprising administering to a mammal in need thereof a therapeutic amount of an agent that decreases the production of a Gob-4 protein.
 - 17. The method of claim 16, wherein said agent that decreases the production of said Gob-4 protein is a nucleic acid.
 - 18. The method of claim 17, wherein said nucleic acid is a ribonucleic acid.
- 19. The method of claim 18, wherein said ribonucleic acid has a nucleotide sequence that is complementary to a portion of the nucleotide sequence set forth in SEQ ID NO:1 or SEQ ID NO:3 encoding said acidic mammalian protein.
 - 20. The method of claim 18, wherein said ribonucleic acid is RNA interference.

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